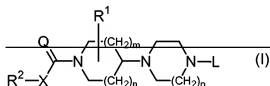


**Amendments to the Claims:**

This listing of claims will replace all prior versions, and listings, of claims in the application:

1. (Currently Amended) A pharmaceutical composition comprising a pharmaceutically acceptable carrier and, as active ingredients, an opioid analgesic and a therapeutically effective amount of a compound selected from the group consisting of ~~according to~~ Formula (I)
- (+)-(B)-trans-4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-piperidinyl]-N-(2,6-dimethylphenyl)-1-piperazine acetamide; and



~~the pharmaceutically acceptable acid or base addition salts thereof, the stereochemically isomeric forms thereof, the N-oxide form thereof, and the prodrugs thereof, wherein~~

n — is 0, 1 or 2;

m — is 1 or 2, provided that if m is 2, then n is 1;

p — is 1 or 2;

=Q — is =O or =NR<sup>3</sup>;

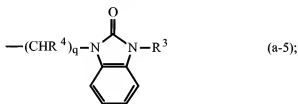
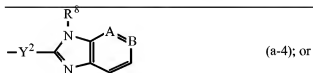
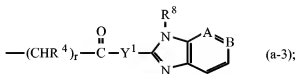
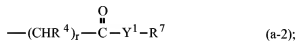
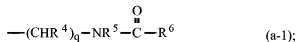
X — is a covalent bond or a bivalent radical of formula —O—, —S—, —NR<sup>3</sup>—;

R<sup>1</sup> — is Ar<sup>1</sup>, Ar<sup>1</sup>C<sub>1-6</sub>alkyl or di(Ar<sup>1</sup>)C<sub>1-6</sub>alkyl, wherein each C<sub>1-6</sub>alkyl group is optionally substituted with hydroxy, C<sub>1-4</sub>alkyloxy, oxo or a ketalized-oxo substituent of formula —O—CH<sub>2</sub>—CH<sub>2</sub>—O— or —O—CH<sub>2</sub>—CH<sub>2</sub>—CH<sub>2</sub>—O—;

R<sup>2</sup> — is Ar<sup>2</sup>, Ar<sup>2</sup>C<sub>1-6</sub>alkyl, Het<sup>1</sup> or Het<sup>1</sup>C<sub>1-6</sub>alkyl;

R<sup>3</sup> — is hydrogen or C<sub>1-6</sub>alkyl;

L — is hydrogen; Ar<sup>2</sup>; C<sub>1-6</sub>alkyl; C<sub>1-6</sub>alkyl substituted with 1 or 2 substituents selected from hydroxy, C<sub>1-6</sub>alkyloxy, Ar<sup>3</sup>, Ar<sup>3</sup>C<sub>1-6</sub>alkyloxy and Het<sup>2</sup>; C<sub>3-6</sub>alkenyl; Ar<sup>3</sup>C<sub>3-6</sub>alkenyl; di(Ar<sup>3</sup>)C<sub>3-6</sub>alkenyl or a radical of formula



wherein

each q — independently is 2, 3 or 4;

each r — is 0, 1, 2, 3 or 4;

each Y<sup>1</sup> — independently is a covalent bond, —O— or NR<sup>3</sup>;

Y<sup>2</sup> — is a covalent bond, C<sub>1-4</sub>alkanediyl or C<sub>1-4</sub>alkylNR<sup>3</sup>;

each A-B — independently is a bivalent radical of formula —CH—CH—, N—CH— or CH=N—;

each R<sup>4</sup> — independently is hydrogen, C<sub>1-6</sub>alkyl, Ar<sup>2</sup> or Ar<sup>2</sup>C<sub>1-6</sub>alkyl;

R<sup>5</sup> — is hydrogen, C<sub>1-6</sub>alkyl or Ar<sup>3</sup>;

R<sup>6</sup> — is C<sub>1-6</sub>alkyl, Ar<sup>3</sup>, Ar<sup>3</sup>C<sub>1-6</sub>alkyl, di(Ar<sup>3</sup>)C<sub>1-6</sub>alkyl, Ar<sup>3</sup>C<sub>3-7</sub>cycloalkyl, or indolyl;

R<sup>7</sup> — is Ar<sup>3</sup>, Ar<sup>3</sup>C<sub>1-6</sub>alkyl; di(Ar<sup>3</sup>)C<sub>1-6</sub>alkyl; C<sub>1-6</sub>alkyl; C<sub>3-7</sub>cycloalkyl; C<sub>3-7</sub>cycloalkyl substituted with Ar<sup>3</sup>; oxazolyl; oxazolyl substituted with halo

or C<sub>1-6</sub>alkyl; thiazolyl; thiazolyl-substituted with halo or C<sub>1-6</sub>alkyl; imidazolyl; imidazolyl-substituted with Ar<sup>3</sup>; C<sub>1-6</sub>alkyl; Ar<sup>3</sup>C<sub>1-6</sub>alkyl or halo; indolyl; indolyl-substituted with C<sub>1-4</sub>alkyl; 2,3,4-trihydroquinolyl; pyrrolidinyl or furanyl;

each R<sup>8</sup> independently is hydrogen, C<sub>1-6</sub>alkyl, C<sub>3-7</sub>cycloalkyl or a radical of formula of formula

-Alk-R<sup>11</sup> (b-1) or

-Alk-Z-R<sup>12</sup> (b-2);

wherein

Alk is C<sub>1-6</sub>alkanediyl;

Z is a bivalent radical of formula -O-, -S- or -NR<sup>2</sup>-;

R<sup>11</sup> is phenyl; phenyl-substituted with 1 or 2 substituents selected from halo, C<sub>1-6</sub>alkyl or C<sub>1-6</sub>alkyloxy; furanyl; furanyl-substituted with 1 or 2 substituents selected from C<sub>1-6</sub>alkyl or hydroxyC<sub>1-6</sub>alkyl; thienyl; thienyl-substituted with 1 or 2 substituents selected from halo or C<sub>1-6</sub>alkyl; oxazolyl; oxazolyl-substituted with 1 or 2 C<sub>1-6</sub>alkyl substituents; thiazolyl; thiazolyl-substituted with 1 or 2 C<sub>1-6</sub>alkyl substituents; pyridinyl or pyridinyl-substituted with 1 or 2 C<sub>1-6</sub>alkyl substituents;

R<sup>12</sup> is C<sub>1-6</sub>alkyl or C<sub>1-6</sub>alkyl-substituted with hydroxy, carboxyl or C<sub>1-6</sub>alkyloxy-carbonyl;

Ar<sup>1</sup> is phenyl; phenyl-substituted with 1, 2 or 3 substituents each independently selected from the group consisting of halo, C<sub>1-4</sub>alkyl, haloC<sub>1-4</sub>alkyl, cyano, aminocarbonyl, C<sub>1-4</sub>alkyloxy and haloC<sub>1-4</sub>alkyloxy;

Ar<sup>2</sup> is naphthalenyl; phenyl; phenyl-substituted with 1, 2 or 3 substituents each independently selected from the group consisting of hydroxy, halo, cyano, nitro, amino, mono- or di(C<sub>1-4</sub>alkyl)amino, C<sub>1-4</sub>alkyl, haloC<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxy, haloC<sub>1-4</sub>alkyloxy, carboxyl, C<sub>1-4</sub>alkyloxy-carbonyl, aminocarbonyl and mono- and di(C<sub>1-4</sub>alkyl)aminocarbonyl;

Ar<sup>3</sup> is phenyl or phenyl-substituted with 1, 2 or 3 substituents selected from the group consisting of halo, hydroxy, amino, nitro, aminocarbonyl, C<sub>1-6</sub>alkyl, haloC<sub>1-6</sub>alkyl and C<sub>1-6</sub>alkyloxy;

Het<sup>1</sup> is a monocyclic heterocycle selected from pyrrolyl, pyrazolyl, imidazolyl, furanyl, thienyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, pyridinyl, pyrimidinyl, pyrazinyl and pyridazinyl; or a bicyclic heterocycle selected from

the group consisting of quinolinyl, quinoxaliny, indolyl, benzimidazolyl, benzoxazolyl, benzisoxazolyl, benzothiazolyl, benzisothiazolyl, benzofuranyl and benzothieryl; each monocyclic and bicyclic heterocycle may optionally be substituted on a carbon atom by 1 or 2 substituents selected from the group consisting of halo, C<sub>1-4</sub>alkyl or mono-, di- and tri(halo)methyl; and

Het<sup>2</sup> is a heterocycle selected from the group consisting of 1,4 dihydro-5-oxo-tetrazol-1-yl, imidazo[1,2-a]pyridinyl, oxazolyl and imidazolyl; each of said heterocycles may be substituted with 1 or where possible 2 substituents selected from the group consisting of C<sub>1-4</sub>alkyl and Ar<sup>2</sup>.

2. (Canceled)

3. (Canceled)

4. (Canceled)

5. (Canceled)

6. (Canceled)

7. (Canceled)

8. (Canceled)

9. (Withdrawn) A pharmaceutical composition according to claim 1 wherein, the pharmaceutical composition comprises a compound selected from the group consisting of:

- 4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-piperidinyl]-N-(2,6-dimethylphenyl)-1-piperazine acetamide;
- 4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-piperidinyl]-N-(1-phenylcyclohexyl)-1-piperazine acetamide;
- 1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-[4-[(1-pyrrolidinylcarbonyl)benzyl]-1-piperazinyl]piperidine;

- 1-[3,5-bis(trifluoromethyl)benzoyl]-4-[4-[1-[(2-methyl-5-oxazolyl)methyl]-1*H*-benzimidazol-2-yl]-1-piperazinyl]-2-(phenylmethyl)piperidine;
  - 4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-[(4-trifluoromethylphenyl)methyl]-4-piperidinyl]-*N*-(2,6-dimethylphenyl)-1-piperazine acetamide; and
  - 4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-[(3,4-dichlorophenyl)methyl]-4-piperidinyl]-*N*-(2,6-dimethylphenyl)-1-piperazine acetamide.
10. (Currently Amended) A pharmaceutical composition according to claim 1 wherein, the pharmaceutical composition comprises ~~a compound selected from the group consisting of:~~
- ~~(+)-(B)-*trans*-4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-piperidinyl]-*N*-(2,6-dimethylphenyl)-1-piperazine acetamide;~~
  - ~~(-)-(B)-*cis*-4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-piperidinyl]-*N*-(2,6-dimethylphenyl)-1-piperazine acetamide; and~~
  - ~~(+)-(B)-*trans*-4-[1-[3,5-bis(trifluoromethyl)benzoyl]-2-(phenylmethyl)-4-piperidinyl]-*N*-(2,6-dimethylphenyl)-1-piperazine acetamide (L)-malic acid (1:1).~~
11. (Previously Amended) A pharmaceutical composition according to claim 1 wherein, the pharmaceutical composition is formulated for simultaneous, separate or sequential use.
12. (Previously Amended) A pharmaceutical composition according to claim 1 wherein, the opioid analgesic is one or more compounds selected from the group consisting of alfentanil, buprenorphine, butorphanol, carfentanyl, codeine, diacetylmorphine, dihydrocodeine, fentanyl, hydrocodone, hydromorphone, levorphanol, lofentanyl, meperidine, methadone, morphine, nalbuphine, oxycodone, oxymorphone, pentazocine, propoxyphene, remifentanyl and sufentanyl; and derivatives and pharmaceutical acceptable salts thereof.
13. (Previously Amended) A pharmaceutical composition according to claim 12 wherein the opioid analgesic is one or more compounds selected from the group consisting of oxycodone, codeine, morphine, fentanyl, buprenorphine, hydrocodone, hydromorphone and pharmaceutical acceptable salts and derivatives thereof.

14. (Previously Amended) A pharmaceutical composition according to claim 1 where, the pharmaceutical composition is in a form suitable to be orally administered.
15. (Canceled)
16. (Canceled)
17. (Canceled)
18. (Canceled)
19. (Canceled)
20. (Canceled)
21. (Withdrawn) A method for treating pain and/or nociception comprising administering to a person in need thereof an effective amount of a pharmaceutical composition according to claim 1.
22. (Withdrawn) A method for treating acute and chronic pain selected from the group consisting of inflammatory, post-operative, emergency room (ER), breakthrough, neuropathic and cancer pain comprising administering to one in need thereof an effective amount of a pharmaceutical composition according to claim 1.
23. (Withdrawn) A method for treating emesis in opioid-based treatments of pain comprising administering to one in need thereof an effective amount of a pharmaceutical composition according to claim 1.
24. (Withdrawn) A method for treating nausea and vomiting in opioid-based treatments of pain comprising administering to one in need thereof an effective amount of a pharmaceutical composition according to claim 23.
25. (Withdrawn) A method for treating respiratory depression in opioid-based treatments of pain comprising administering to one in need thereof an effective amount of an

NK<sub>1</sub>-receptor antagonist selected from an NK<sub>1</sub>-receptor antagonist according to Formula (I), the pharmaceutically acceptable acid or base addition salts thereof, the stereochemically isomeric forms thereof, the *N*-oxide form thereof and prodrugs thereof.

26. (Withdrawn) A method for reducing and/or overcoming the tolerance observed with opioids in opioid-based treatments of pain comprising administering to one in need thereof an effective amount of an NK<sub>1</sub>-receptor antagonist selected from the group consisting of an NK<sub>1</sub>-receptor antagonist according to Formula (I), the pharmaceutically acceptable acid or base addition salts thereof, the stereochemically isomeric forms thereof, the *N*-oxide form thereof and prodrugs thereof.